WHAT IS CLAIMED IS:

1. A compound of Formula I:

$$(R^3)_{\overline{n}} \xrightarrow{\overline{y}}_{Z} N \xrightarrow{N} R^1$$

$$I \qquad (R^2)_{\overline{p}}$$

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or a pharmaceutically acceptable salt or stereoisomer thereof, wherein

w, x, y and z are independently selected from CH, CH_2 and N, provided that at the most only one of w, x, y and z is N and one of w, x, y and z is N only when both dashed lines represent a double bond;

a dashed line represents an optional double bond;

a is 0 or 1;

15 b is 0 or 1;

m is 0, 1, or 2;

n is 0 to 2;

p is 1 to 3;

r is 0 or 1;

20 s is 0 or 1;

R¹ is selected from:

- 1) H,
- 2) C_1 - C_{10} alkyl,
- 25 3) aryl,
 - 4) C_2 - C_{10} alkenyl,
 - 5) C2-C₁₀ alkynyl,
 - 6) C₁-C₆ perfluoroalkyl,
 - 7) C₁-C₆ aralkyl,
- 30 8) C3-C8 cycloalkyl, and

9) heterocyclyl,

said alkyl, aryl, alkenyl, alkynyl, cycloalkyl, aralkyl and heterocyclyl is optionally substituted with one or more substituents selected from R⁴;

- 5 R² and R³ is independently selected from:
 - 1) $(C=O)_aO_bC_1-C_{10}$ alkyl,
 - 2) (C=O)_aO_baryl,
 - 3) $(C=O)_aO_bC_2-C_{10}$ alkenyl,
 - 4) $(C=O)_aO_bC_2-C_{10}$ alkynyl,
- 10 5) CO₂H,
 - 6) halo,
 - 7) OH,
 - 8) ObC1-C6 perfluoroalkyl,
 - 9) $(C=O)_aNR^6R^7$,
- 15 10) CN,
 - 11) (C=O)_aO_bC₃-C₈ cycloalkyl,
 - 12) (C=O)_aO_bheterocyclyl,
 - 13) $SO_2NR^6R^7$, and
 - 14) SO_2C_1 - C_{10} alkyl,
- said alkyl, aryl, alkenyl, alkynyl, cycloalkyl, and heterocyclyl is optionally substituted with one or more substituents selected from R⁴;

R⁴ is independently selected from:

- 1) $(C=O)_aO_bC_1-C_{10}$ alkyl,
- 25 2) (C=O)_aO_baryl,
 - 3) C2-C₁₀ alkenyl,
 - 4) C2-C₁₀ alkynyl,
 - 5) (C=O)_aO_b heterocyclyl,
 - 6) CO₂H,
- 30 7) halo,
 - 8) CN,
 - 9) OH,
 - 10) ObC1-C6 perfluoroalkyl,
 - 11) $O_a(C=O)_bNR^6R^7$,
- 35 12) oxo,

- 13) CHO,
- $(N=0)R^6R^7$, or
- 15) (C=O)_aO_bC₃-C₈ cycloalkyl,
- 16) SO₂C₁-C₁₀alkyl,
- 17) $SO_2NR^6R^7$,

said alkyl, aryl, alkenyl, alkynyl, heterocyclyl, and cycloalkyl optionally substituted with one or more substituents selected from R⁵;

R⁵ is selected from:

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- 10 1) $(C=O)_{r}O_{s}(C_{1}-C_{10})$ alkyl,
 - 2) $O_r(C_1-C_3)$ perfluoroalkyl,
 - 3) (C_0-C_6) alkylene- $S(O)_mR^a$,
 - 4) oxo,
 - 5) OH,
- 15 6) halo,
 - 7) CN,
 - 8) $(C=O)_rO_s(C_2-C_{10})$ alkenyl,
 - 9) $(C=O)_rO_s(C_2-C_{10})$ alkynyl,
 - 10) $(C=O)_{\Gamma}O_{S}(C_{3}-C_{6})$ cycloalkyl,
- 20 $(C=O)_rO_s(C_0-C_6)$ alkylene-aryl,
 - 12) $(C=O)_rO_s(C_0-C_6)$ alkylene-heterocyclyl,
 - 13) $(C=O)_rO_s(C_0-C_6)$ alkylene- $N(R^b)_2$,
 - $C(O)R^a$,
 - 15) (C₀-C₆)alkylene-CO₂R^a
- 25 16) C(O)H,

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- 17) (C₀-C₆)alkylene-CO₂H, and
- 18) $C(O)N(R^b)_2$,

said alkyl, alkenyl, alkynyl, cycloalkyl, aryl, and heterocyclyl is optionally substituted with up to three substituents selected from R^b, OH, (C₁-C₆)alkoxy, halogen, CO₂H, CN, O(C=O)C₁-C₆ alkyl, oxo, and N(R^b)₂;

R6 and R7 are independently selected from:

- 1) H,
- 2) (C=O)O_bC₁-C₁₀ alkyl,
- 35 (C=O)O_bC₃-C₈ cycloalkyl,

- 4) (C=O)Obaryl,
- 5) (C=O)Obheterocyclyl,
- 6) C_1 - C_{10} alkyl,
- 7) aryl,

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- 8) C2-C₁₀ alkenyl,
- 9) C_2 - C_{10} alkynyl,
- 10) heterocyclyl,
- 11) C3-C8 cycloalkyl,
- 12) SO₂Ra, and
- 10 13) $(C=O)NRb_2$,

said alkyl, cycloalkyl, aryl, heterocylyl, alkenyl, and alkynyl is optionally substituted with one or more substituents selected from R⁵, or

R⁶ and R⁷ can be taken together with the nitrogen to which they are attached to form a monocyclic or bicyclic heterocycle with 4-7 members in each ring and optionally containing, in addition to the nitrogen, one or two additional heteroatoms selected from N, O and S, said monocyclic or bicyclic heterocycle optionally substituted with one or more substituents selected from R⁵;

 R^a is (C1-C6)alkyl, (C3-C6)cycloalkyl, aryl, or heterocyclyl; and

Rb is H, (C1-C6)alkyl, (C1-C6)alkyl-NRa2, (C1-C6)alkyl-NH2, (C1-C6)alkyl-NHRa, aryl, heterocyclyl, (C3-C6)cycloalkyl, (C=O)OC1-C6 alkyl, (C=O)C1-C6 alkyl or S(O)2Ra.

2. The compound according to Claim 1 of the formula II:

wherein a, w, x, y, z, dashed line, R^3 , R^4 , R^6 and R^7 are defined as in Claim 1 for the compound of the Formula I; and

n is 0 or 1;

p' is 0 to 2;

R² is selected from:

5 1) $(C=O)_aC_1-C_{10}$ alkyl,

- 2) (C=O)_aaryl,
- 3) $(C=O)_aNR^6R^7$,
- 4) (C=O)_aC₃-C₈ cycloalkyl,
- 5) (C=O)_aheterocyclyl,
- 6) $SO_2NR^6R^7$, and
 - 7) SO_2C_1 - C_{10} alkyl,

said alkyl, aryl, cycloalkyl, and heterocyclyl is optionally substituted with one or more substituents selected from R⁴;

15 R^{2a} is selected from: halogen and (C₁-C₆)alkyl; and

R^{4a} and R^{4b} are independently selected from: hydrogen, halogen and (C₁-C₆)alkyl, provided that at lease one is not hydrogen, or

- 20 R^{4a} and R^{4b} are combined to form a diradical selected from –CH₂CH₂CH₂CH₂-, –CH₂CH₂CH₂-, –CH₂CH₂-, –CH₂CH₂-, –CH₂CH₂-, –CH₂CH₂-, –CH₂-CH₂-, –CH₂-, –CH₂-,
 - 3. A compound of the formula III, or a pharmaceutically acceptable salt or stereoisomer thereof,

$$\mathbb{R}^{3a}$$
 \mathbb{R}^{3b}
 \mathbb{R}^{4a}
 \mathbb{R}^{2a}
 \mathbb{R}^{2a}
 \mathbb{R}^{2b}

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wherein

b is 0 or 1;

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m is 0, 1 or 2;
         p' is 0 to 2;
         r is
                    0 or 1;
                    0 \text{ or } 1;
         s is
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         R<sup>2</sup> is (C<sub>1</sub>-C<sub>6</sub>)alkylene-NR<sup>6</sup>R<sup>7</sup>; said alkylene is optionally substituted with up to three substituents
         selected from OH, (C1-C6)alkoxy, halogen, CO2H, CN, O(C=O)C1-C6 alkyl, oxo, and NR<sup>6</sup>R<sup>7</sup>;
         R<sup>2a</sup> is selected from: halogen and (C<sub>1</sub>-C<sub>6</sub>)alkyl;
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         R<sup>3a</sup> and R<sup>3b</sup> are independently selected from: hydrogen and halogen; and
         R<sup>4a</sup> and R<sup>4b</sup> are independently selected from: hydrogen, halogen, and (C<sub>1</sub>-C<sub>6</sub>)alkyl, provided that at
         least one is not hydrogen;
15
         R<sup>5</sup> is selected from:
                    1)
                               (C=O)_rO_s(C_1-C_{10})alkyl,
                    2)
                                O<sub>r</sub>(C<sub>1</sub>-C<sub>3</sub>)perfluoroalkyl,
                               (C<sub>0</sub>-C<sub>6</sub>)alkylene-S(O)<sub>m</sub>Ra,
                    3)
                    4)
                                oxo,
20
                    5)
                               OH,
                    6)
                               halo,
                    7)
                               CN,
                    8)
                               (C=O)rOs(C2-C10)alkenyl,
                               (C=O)rOs(C2-C10)alkynyl,
25
                    9)
                               (C=O)<sub>r</sub>O<sub>s</sub>(C3-C6)cycloalkyl,
                    10)
                               (C=O)rOs(C0-C6)alkylene-aryl,
                    11)
                               (C=O)<sub>r</sub>O<sub>s</sub>(C<sub>0</sub>-C<sub>6</sub>)alkylene-heterocyclyl,
                    12)
                               (C=O)_rO_s(C_0-C_6)alkylene-N(R^b)_2,
                    13)
                               C(O)R^{a}
                    14)
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                    15)
                               (C<sub>0</sub>-C<sub>6</sub>)alkylene-CO<sub>2</sub>R<sup>a</sup>.
                               C(O)H,
                    16)
                               (C<sub>0</sub>-C<sub>6</sub>)alkylene-CO<sub>2</sub>H, and
                    17)
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 $C(O)N(R^b)_2$,

18)

said alkyl, alkenyl, alkynyl, cycloalkyl, aryl, and heterocyclyl is optionally substituted with up to three substituents selected from R^b , OH, (C_1-C_6) alkoxy, halogen, CO_2H , CN, $O(C=O)C_1-C_6$ alkyl, oxo, and $N(R^b)_2$;

- 5 R6 and R7 are independently selected from:
 - 1) H,
 - 2) $(C=O)O_bC_1-C_{10}$ alkyl,
 - 3) (C=O)ObC3-C8 cycloalkyl,
 - 4) (C=O)Obaryl,
- 10 5) (C=O)Obheterocyclyl,
 - 6) C₁-C₁₀ alkyl,
 - 7) aryl,
 - 8) C2-C₁₀ alkenyl,
 - 9) C₂-C₁₀ alkynyl,
- 15 10) heterocyclyl,
 - 11) C3-C8 cycloalkyl,
 - 12) SO₂Ra, and
 - 13) $(C=O)NRb_2$,

said alkyl, cycloalkyl, aryl, heterocylyl, alkenyl, and alkynyl is optionally substituted with one or more substituents selected from R⁵, or

R⁶ and R⁷ can be taken together with the nitrogen to which they are attached to form a monocyclic or bicyclic heterocycle with 4-7 members in each ring and optionally containing, in addition to the nitrogen, one or two additional heteroatoms selected from N, O and S, said monocyclic or bicyclic heterocycle optionally substituted with one or more substituents selected from R⁵;

- R^a is (C1-C6)alkyl, (C3-C6)cycloalkyl, aryl, or heterocyclyl; and
- R^b is H, (C₁-C₆)alkyl, (C₁-C₆)alkyl-NR^a₂, (C₁-C₆)alkyl-NH₂, (C₁-C₆)alkyl-NHR^a, aryl, heterocyclyl, (C₃-C₆)cycloalkyl, (C=O)OC₁-C₆ alkyl, (C=O)C₁-C₆ alkyl or S(O)₂R^a.
 - 4. The compound according to Claim 3, or the pharmaceutically acceptable salt or stereoisomer thereof, wherein p', R^{2a}, R^{3a}, R^{3b}, R^{4a}, R^{4b} and R⁵ are as defined for Formula III in Claim 3 and

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R² is (C₁-C₆)alkylene-NR⁶R⁷;

R⁶ and R⁷ are independently selected from:

1) H,

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- C_1 - C_{10} alkyl,
- 3) aryl,
- 4) heterocyclyl,
- 5) C2-C₁₀ alkenyl,
- 6) C2-C₁₀ alkynyl, and
- 10 7) C3-C8 cycloalkyl,

said alkyl, cycloalkyl, aryl, heterocylyl, alkenyl, and alkynyl is optionally substituted with one or more substituents selected from R⁵, or

R⁶ and R⁷ can be taken together with the nitrogen to which they are attached to form a monocyclic or bicyclic heterocycle with 4-7 members in each ring and optionally containing, in addition to the nitrogen, one or two additional heteroatoms selected from N, O and S, said monocyclic or bicyclic heterocycle optionally substituted with one or more substituents selected from R⁵.

5. A compound selected from:

20 2-(2-bromophenyl)-3-(4-methylphenyl)quinazolin-4(3H)-one;

2-(2-bromophenyl)-3-(4-methylphenyl)-quinazolin-4(3H)-one;

25 2-(2-chlorophenyl)-3-(4-methylphenyl)-quinazolin-4(3H)-one;

2-(2,4-dichlorophenyl)-3-(4-methylphenyl)quinazolin-4(3H)-one;

2-(2-bromophenyl)-3-(4-chlorophenyl)-quinazolin-4(3H)-one;

2-(2-bromophenyl)-3-(3-fluoro-4-methylphenyl)-quinazolin-4(3H)-one;

3-(3a,7a-dihydro-1H-indol-5-yl)-2-(2-bromophenyl)-quinazolin-4(3H)-one;

35 6-chloro-2-(2-chlorophenyl)-3-(3-fluoro-4-methylphenyl)-quinazolin-4(3H)-one;

2-(2-chlorophenyl)-3-(3-fluoro-4-methylphenyl)quinazolin-4(3H)-one;

2-(2-methylphenyl)-3-(4-methylphenyl)-quinazolin-4(3H)-one;

7-chloro-2-(2-chlorophenyl)-3-(3-fluoro-4-methylphenyl)quinazolin-4(3H)-one;

- 2-(2-bromophenyl)-7-chloro-3-(3-fluoro-4-methylphenyl)quinazolin-4(3H)-one;
- 7-chloro-2-(2-chlorophenyl)-3-(1H-indol-5-yl)quinazolin-4(3H)-one;

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- 5 2-(2-bromophenyl)-7-chloro-3-(4-chloro-3-fluorophenyl)quinazolin-4(3H)-one;
 - 2-(2-bromophenyl)-3-(3-fluoro-4-methyl-phenyl)pyrido[2,3-d]pyrimidin-4(3H)-one;
 - 2-(5-bromo-2-chlorophenyl)-7-chloro-3-(3-fluoro-4-methylphenyl)quinazolin-4(3H)-one;
 - 2-(4-bromo-2-chlorophenyl)-7-chloro-3-(3-fluoro-4-methylphenyl)quinazolin-4(3H)-one;
 - 2-(2-chlorophenyl)-3-(3-fluoro-4-methylphenyl)-5,6,7,8-tetrahydroquinazolin-4(3H)-one;
- 7-chloro-2-{2-chloro-3-[(dimethylamino)methyl]phenyl}-3-(4-chloro-3-fluorophenyl)quinazolin-4(3H)-one;
 - $7-chloro-3-(4-chloro-3-fluorophenyl)-2-\{2-chloro-5-[(4-methylpiperazin-1-yl)methyl]phenyl\} quinazolin-4(3H)-one;$
 - 7-chloro-3-(4-chloro-3-fluorophenyl)-2-{2-chloro-3-[(methylamino)methyl]-phenyl}quinazolin-4(3H)-one;
- 7-chloro-3-(4-chloro-3-fluorophenyl)-2-{2-chloro-3-[(4-methylpiperazin-1-yl)methyl]phenyl}quinazolin-25 4(3H)-one;
 - 7-chloro-2-{2-chloro-3-[(ethylamino)methyl]phenyl}-3-(4-chloro-3-fluorophenyl)quinazolin-4(3H)-one;
- 7-chloro-3-(4-chloro-3-fluorophenyl)-2-{2-chloro-3-[(isopropylamino)methyl]-phenyl}quinazolin-4(3H)-30 one;
 - 7-chloro-2-{2-chloro-3-[(cyclobutylamino)methyl]phenyl}-3-(4-chloro-3-fluorophenyl)quinazolin-4(3H)-one;
- 2-[3-(azetidin-1-ylmethyl)-2-chlorophenyl]-7-chloro-3-(4-chloro-3-fluorophenyl)quinazolin-4(3H)-one;
 7-chloro-3-(4-chloro-3-fluorophenyl)-2-[2-chloro-3-(pyrrolidin-1-ylmethyl)phenyl]quinazolin-4(3H)-one;

7-chloro-3-(4-chloro-3-fluorophenyl)-2-(2-chloro-3-{[(3S)-3-hydroxypyrrolidin-1-yl]methyl}phenyl)quinazolin-4(3H)-one;

- 7-chloro-3-(4-chloro-3-fluorophenyl)-2-(2-chloro-3-{[(3S)-3-(methoxymethyl)pyrrolidin-1-yl]methyl}phenyl)quinazolin-4(3H)-one;
 - $7-chloro-3-(4-chloro-3-fluorophenyl)-2-\{2-chloro-3-[(pyrrolidin-3-ylamino)methyl]phenyl\} quinazolin-4(3H)-one;$
- 7-chloro-3-(4-chloro-3-fluorophenyl)-2-[2-chloro-3-(morpholin-4-ylmethyl)phenyl]quinazolin-4(3H)-one;
 - 7-chloro-3-(4-chloro-3-fluorophenyl)-2-[2-chloro-3-(piperidin-1-ylmethyl)phenyl]quinazolin-4(3H)-one;
- 2-{3-[(4-aminopiperidin-1-yl)methyl]-2-chlorophenyl}-7-chloro-3-(4-chloro-3-fluorophenyl)quinazolin-4(3H)-one;
 - 7-chloro-3-(4-chloro-3-fluorophenyl)-2-{2-chloro-3-[(piperidin-4-ylamino)methyl]phenyl}quinazolin-4(3H)-one;
 - $7-chloro-3-(4-chloro-3-fluorophenyl)-2-\{2-chloro-3-[(4-fluoropiperidin-1-yl)methyl]phenyl\} quinazolin-4(3H)-one;$
 - 7-chloro-3-(4-chloro-3-fluorophenyl)-2-[2-chloro-3-(piperazin-1-ylmethyl)phenyl]quinazolin-4(3H)-one;
 - $2-\{3-[(4-acetylpiperazin-1-yl)methyl]-2-chlorophenyl\}-7-chloro-3-(4-chloro-3-fluorophenyl)quinazolin-4(3H)-one;$
- 7-chloro-3-(4-chloro-3-fluorophenyl)-2-(2-chloro-3-{[4-(methylsulfonyl)piperazin-1-yl]methyl}phenyl)quinazolin-4(3H)-one;

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7-chloro-3-(4-chloro-3-fluorophenyl)-2-(2-chloro-3-{[(2-hydroxyethyl)amino]-methyl}phenyl)quinazolin-4(3H)-one;

 $\label{lem:condition} $$7$-chloro-2-[2-chloro-3-({[2-(dimethylamino)ethyl]amino}]methyl)$ phenyl]-3-(4-chloro-3-fluorophenyl)$ quinazolin-4(3H)-one;$

- 7-chloro-3-(4-chloro-3-fluorophenyl)-2-(2-chloro-3-{[(2-morpholin-4-
- 5 ylethyl)amino]methyl}phenyl)quinazolin-4(3H)-one;

- $2-\{3-[(3-aminopyrrolidin-1-yl)methyl]-2-chlorophenyl\}-7-chloro-3-(4-chloro-3-fluorophenyl)quinazolin-4(3H)-one;$
- 7-chloro-3-(4-chloro-3-fluorophenyl)-2-[2-chloro-3-({[(1-methylpiperidin-3-yl)methyl]amino}methyl)phenyl]quinazolin-4(3H)-one;
 - $2-(3-\{[3-(aminomethyl)-1-methyl-1lambda~5~-piperidin-1-yl]methyl\}-2-chlorophenyl)-7-chloro-3-(4-chloro-3-fluorophenyl)quinazolin-4(3H)-one;$
- 2-{3-[(benzylamino)methyl]-2-chlorophenyl}-7-chloro-3-(4-chloro-3-fluorophenyl)quinazolin-4(3H)-one;
- 7-chloro-3-(4-chloro-3-fluorophenyl)-2-{2-chloro-5-[(4-methylpiperazin-1-yl)methyl]phenyl}quinazolin-20 4(3H)-one;
 - $7-chloro-2-\{2-chloro-5-[(ethylamino)methyl]phenyl\}-3-(4-chloro-3-fluorophenyl)quinazolin-4(3H)-one;\\$
- 7-chloro-3-(4-chloro-3-fluorophenyl)-2-{2-chloro-5-[(isopropylamino)methyl]-phenyl}quinazolin-4(3H)-25 one;
 - 7-chloro-3-(4-chloro-3-fluorophenyl)-2-[2-chloro-5-(pyrrolidin-1-ylmethyl)phenyl] quinazolin-4(3H)-one;
- 7-chloro-3-(4-chloro-3-fluorophenyl)-2-{2-chloro-5-[(pyrrolidin-3-ylamino)methyl]phenyl}quinazolin-30 4(3H)-one;
 - 7-chloro-3-(4-chloro-3-fluorophenyl)-2-[2-chloro-5-(morpholin-4-ylmethyl)phenyl]quinazolin-4(3H)-one;
- 35 7-chloro-3-(4-chloro-3-fluorophenyl)-2-[2-chloro-5-(piperidin-1-ylmethyl)phenyl]quinazolin-4(3H)-one;

 $7-chloro-3-(4-chloro-3-fluorophenyl)-2-\{2-chloro-5-[(piperidin-4-ylamino)methyl]phenyl\} quinazolin-4(3H)-one;$

- 5 7-chloro-3-(4-chloro-3-fluorophenyl)-2-[2-chloro-5-(piperazin-1-ylmethyl)phenyl]quinazolin-4(3H)-one;
 - 7-chloro-3-(4-chloro-3-fluorophenyl)-2-(2-chloro-5-{[4-(methylsulfonyl)piperazin-1-yl]methyl}phenyl)quinazolin-4(3H)-one; and
- 7-chloro-2-[2-chloro-5-({[2-(dimethylamino)ethyl]amino}methyl)phenyl]-3-(4-chloro-3-fluorophenyl)quinazolin-4(3H)-one;

or a pharmaceutically acceptable salt thereof.

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- 6. A pharmaceutical composition that is comprised of a compound in accordance with Claim 1 and a pharmaceutically acceptable carrier.
 - 7. A pharmaceutical composition that is comprised of a compound in accordance with Claim 3 and a pharmaceutically acceptable carrier.
 - 8. A method of treating or preventing cancer in a mammal in need of such treatment that is comprised of administering to said mammal a therapeutically effective amount of a compound of Claim 1.
- 9. A method of treating or preventing cancer in a mammal in need of such treatment that is comprised of administering to said mammal a therapeutically effective amount of a compound of Claim 3.
- 10. A method of treating cancer or preventing cancer in accordance with Claim 8
 wherein the cancer is selected from cancers of the brain, genitourinary tract, lymphatic system, stomach, larynx and lung.
 - 11. A method of treating or preventing cancer in accordance with Claim 8 wherein the cancer is selected from histiocytic lymphoma, lung adenocarcinoma, small cell lung cancers, pancreatic cancer, glioblastomas and breast carcinoma.

12. A process for making a pharmaceutical composition which comprises combining a compound of Claim 1 with a pharmaceutically acceptable carrier.

- 5 13. The composition of Claim 6 further comprising a second compound selected from: an estrogen receptor modulator, an androgen receptor modulator, a retinoid receptor modulator, a cytotoxic/cytostatic agent, an antiproliferative agent, a prenyl-protein transferase inhibitor, an HMG-CoA reductase inhibitor, an HIV protease inhibitor, a reverse transcriptase inhibitor, an angiogenesis inhibitor, a PPAR-γ agonist, a PPAR-δ agonist; an inhibitor of cell proliferation and survival signaling, an agent that interfers with a cell cycle checkpoint, and an apoptosis inducing agent.
 - 14. The composition of Claim 13, wherein the second compound is an angiogenesis inhibitor selected from the group consisting of a tyrosine kinase inhibitor, an inhibitor of epidermal-derived growth factor, an inhibitor of fibroblast-derived growth factor, an inhibitor of platelet derived growth factor, an MMP inhibitor, an integrin blocker, interferon-α, interleukin-12, pentosan polysulfate, a cyclooxygenase inhibitor, carboxyamidotriazole, combretastatin A-4, squalamine, 6-O-(chloroacetyl-carbonyl)-fumagillol, thalidomide, angiostatin, troponin-1, and an antibody to VEGF.
- The composition according to Claim 13 further comprising a proteosome inhibitor.

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- 16. The composition according to Claim 13 further comprising a aurora kinase inhibitor.
- 25 The composition according to Claim 13 further comprising a Raf kinase inhibitor.
 - 18. The composition according to Claim 13 further comprising a serine/threonine kinase inhibitor.
 - 19. The composition according to Claim 13 further comprising an inhibitor of another mitotic kinesin which is not KSP.
- The composition of Claim 13, wherein the second compound is an estrogen receptor modulator selected from tamoxifen and raloxifene.

21. A method of treating cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 in combination with radiation therapy.

- 5 22. A method of treating or preventing cancer that comprises administering a therapeutically effective amount of a compound of Claim 1 in combination with a compound selected from: an estrogen receptor modulator, an androgen receptor modulator, retinoid receptor modulator, a cytotoxic/cytostatic agent, an antiproliferative agent, a prenyl-protein transferase inhibitor, an HMG-CoA reductase inhibitor, an HIV protease inhibitor, a reverse transcriptase inhibitor, an angiogenesis inhibitor, a PPAR-γ agonist, a PPAR-δ agonist, an inhibitor of inherent multidrug resistance, an anti-emetic agent, an agent useful in the treatment of anemia, an agent useful in the treatment of neutropenia, an immunologic-enhancing drug, an inhibitor of cell proliferation and survival signaling, an agent that interfers with a cell cycle checkpoint, and an apoptosis inducing agent.
- 23. A method of treating cancer that comprises administering a therapeutically effective amount of a compound of Claim 1 in combination with radiation therapy and a compound selected from: an estrogen receptor modulator, an androgen receptor modulator, retinoid receptor modulator, a cytotoxic/cytostatic agent, an antiproliferative agent, a prenyl-protein transferase inhibitor, an HMG-CoA reductase inhibitor, an HIV protease inhibitor, a reverse transcriptase inhibitor, an angiogenesis inhibitor, a PPAR-γ agonist, a PPAR-δ agonist, an inhibitor of inherent multidrug resistance, an anti-emetic agent, an agent useful in the treatment of anemia, an agent useful in the treatment of neutropenia, an immunologic-enhancing drug, an inhibitor of cell proliferation and survival signaling, an agent that interfers with a cell cycle checkpoint, and an apoptosis inducing agent.
- 24. A method of treating or preventing cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 and paclitaxel or trastuzumab.
 - 25. A method of treating or preventing cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 and a GPIIb/IIIa antagonist.
 - 26. The method of Claim 25 wherein the GPIIb/IIIa antagonist is tirofiban.
 - 27. A method of treating or preventing cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 in combination with a COX-2 inhibitor.

28. A method of treating or preventing cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 in combination with a proteosome inhibitor.

- 29. A method of treating or preventing cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 in combination with an aurora kinase inhibitor.
 - 30. A method of treating or preventing cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 in combination with a Raf kinase inhibitor.
 - 31. A method of treating or preventing cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 in combination with a serine/threonine kinase inhibitor.
- 15 32. A method of treating or preventing cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 in combination with an inhibitor of a mitotic kinesin that is not KSP.
- 33. A method of modulating mitotic spindle formation which comprises administering a therapeutically effective amount of a compound of Claim 1.

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34. A method of inhibiting the mitotic kinesin KSP which comprises administering a therapeutically effective amount of a compound of Claim 1.